

Claims

1. Nucleic acid which codes for the α chain of a human T cell receptor, or for a functional derivative or a fragment thereof and which comprises a CDR3 region formed from a combination of a Va20 and Ja22 gene segment.

2. Nucleic acid which codes for the α chain of a human T cell receptor, or for a functional derivative or a fragment thereof and comprises a CDR3 region selected from:

- (a) a nucleotide sequence coding for the amino acid sequence (SEQ ID NO:23)

Y C L (X₁...X_n) S A R Q L T F (I)

in which X₁ ... X_n represents a sequence of 3-5 amino acids,

- (b) a nucleotide sequence which codes for an amino acid sequence which is at least 80 % identical with the amino acid sequence from (a), or
- (c) a nucleotide sequence which codes for an amino acid sequence with an equivalent recognition specificity for the peptide component of the T cell receptor ligands.
- W4 + d59p

- I
3. Nucleic acid as claimed in claim 2,
w h e r e i n
the amino acid sequence $X_1 \dots X_n$ is selected from
the group comprising the amino acid sequences VGG,
VLSG, ATG, VSG, DSG, VVSG, ALAG, APSG and VGR.
- 5d8
F2
sub
F2
4. Nucleic acid as claimed in claim 3,
w h e r e i n
the amino acid sequence $X_1 \dots X_n$ is selected from
the group comprising amino acid sequences VGG, VLSG
and ATG.
5. Vector,
w h e r e i n
it contains at least one copy of a nucleic acid as
claimed in one of the claims 1 to 4.
6. Cell,
w h e r e i n
it expresses a nucleic acid as claimed in one of
the claims 1 to 4.
7. Cell,
w h e r e i n
it is transformed with a nucleic acid as claimed in
one of the claims 1 to 4 or with a vector as
claimed in claim 5.
- II
8. Polypeptide,
w h e r e i n
it is coded by a nucleic acid as claimed in one of
the claims 1 to 4.
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II 9. Polypeptide as claimed in claim 8,
wherein
it comprises the variable domain of the α chain of
a human T cell receptor.

III 10. Nucleic acid which codes for the β chain of a human
T cell receptor, or for a functional derivative or
a fragment thereof and comprises a CDR3 region
formed from a combination of a V β 22 gene segment, a
D β 1 or D β 2 gene segment and a J β gene segment in
particular a J β 2.1, J β 2.3 or J β 2.7 gene segment.

11. Nucleic acid which codes for the β chain of a human
T cell receptor, or for a functional derivative or
a fragment thereof and comprises a CDR3 region
which is selected from:

(a) a nucleotide sequence coding for the amino acid
sequence *(SEQ ID NO: 24) and (SEQ ID NO: 45)*
respectively,

C A (X'₁ ... X'_n) Y/D E Q Y F (II)

in which X'₁ ... X'_n represents a sequence of
5-7 amino acids,

(b) a nucleotide sequence coding for the amino acid
sequence *(SEQ ID NO: 25)*

C A (X''₁ ... X''_n) N E Q F F (III)

in which X''₁ ... X''_n represents a sequence of
5-7 amino acids,

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$$C A (X''_1 \dots X''_n) D T Q Y F \quad (IV)$$

in which $X''_1 \dots X''_n$ represents a sequence of 5-7 amino acids.

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ino acid se
, or

12. Nucleic acid as claimed in claim 11,
w h e r e i n

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13. Nucleic acid as claimed in claim 12,
w h e r e i n

✓

14. Nucleic acid as claimed in claim 11,
w h e r e i n
the amino acid sequence X''₁ ... X''_n represents
SSGTSSY or SSDQGH, or the amino acid sequence
X'''₁ ... X'''_n represents SADSFK
(SEQ ID No. 42) (SEQ ID No. 43) (SEQ ID No. 44)
15. Vector,
w h e r e i n
it contains at least one copy of a nucleic acid as
claimed in one of the claims 10 to 14.
16. Cell,
w h e r e i n
it expresses a nucleic acid as claimed in one of
the claims 10 to 14.
17. Cell,
w h e r e i n
it is transformed with a nucleic acid as claimed in
one of the claims 10 to 14 or with a vector as
claimed in claim 15.
18. Polypeptide,
w h e r e i n
it codes for a nucleic acid as claimed in one of
the claims 10 to 14.
19. Polypeptide as claimed in claim 18,
w h e r e i n
it comprises the variable domain of the β chain of
a human T cell receptor.

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26. Pharmaceutical composition which contains as active component a nucleic acid as claimed in one of the claims 1 to 4 or 10 to 14, a polypeptide as claimed in one of the claims 8, 9 or 18 to 23, a peptide ligand against the polypeptide, an antibody as claimed in claim 23 or 24 or a cell as claimed in claim 6, 7, 16, 17 or 25 optionally together with other active components as well as common pharmaceutical auxiliary agents, additives or carrier substances.

27. Use of a pharmaceutical composition as claimed in claim 26 for the production of an agent for the diagnosis of tumour diseases or a predisposition for a tumour disease.

28. Use of a pharmaceutical composition as claimed in claim 26 for the production of an agent for monitoring the course of the disease in a tumour disease.

29. Use as claimed in claim 27 or 28, *improved MDC*
wherein the detection of T cells that express a polypeptide as claimed in claim 20 as the T cell receptor is carried out in a sample liquid by a nucleic acid hybridization assay, an immunoassay, a test for the binding of specific peptide ligands or a specific T cell activity test.

30. Use of a pharmaceutical composition as claimed in claim 26 for the production of an agent for the prevention or therapy of a tumour disease.

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- VI*
31. Use as claimed in claim 30, *wherein* *wherein* *MDC*
wherein
the agent is suitable for the stimulation of the
growth of T cells that express a polypeptide as
claimed in claim 20 as a T cell receptor.
32. Use as claimed in claim 31,
wherein
the agent is suitable for growth stimulation of the
T cells in vivo.
33. Use as claimed in claim 31 or 32,
wherein
the agent for growth stimulation comprises the
peptide ligand of the T cell receptor or/and the
entire molecule from which the peptide ligand is
derived or a fragment thereof.
34. Use as claimed in claim 31 or 32,
wherein
the growth stimulation includes an antibody that
specifically activates the T cell receptor.
35. Use as claimed in claim 31,
wherein
the growth stimulation is carried out by isolating
specific T cells, in vitro expansion and subsequent
administration of expanded T cells.
36. Use as claimed in one of the claims 27 to 35,
wherein
the tumour disease is a kidney cell carcinoma.

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37. Process for the isolation of T cells that express a polypeptide as claimed in claim 20 as a T cell receptor,
w h e r e i n
a sample containing T cells is contacted with an agent that binds specifically to the CDR3 region of the T cell receptor. T cells that react with the agent are identified and optionally separated from other T cells.
38. Process as claimed in claim 37,
w h e r e i n
the agent is selected from the peptide ligand of T cells, a MHC peptide complex containing the peptide ligand or/and an anti-TCR antibody.
39. Process as claimed in claim 37 or 38 additionally comprising an in vitro expansion of T cells.
40. Process for the isolation of T cells which express a polypeptide as claimed in claim 20 as the T cell receptor,
w h e r e i n
nucleic acid sequences that code for the T cell receptor are introduced into a T cell line and are made to express therein.
41. Process for the isolation of T cells that express a polypeptide as claimed in claim 20 as the T cell receptor,
w h e r e i n
nucleic acid sequences which code for the T cell

receptor are introduced into the germ line of an animal and the T cells are isolated from the resulting transgenic animal or descendants thereof.

42. Transgenic animal,
w h e r e i n
it expresses a polypeptide as claimed in claim 20
as the T cell receptor.
43. Method for the identification of peptide ligands of
a T cell receptor as claimed in claim 20 comprising
the steps:
- (a) isolating RNA from tumour tissue,
 - (b) converting the RNA into double-stranded cDNA
molecules,
 - (c) introducing the cDNA molecules into host cells
to obtain a cDNA bank,
 - (d) transfecting eukaryotic recipient cells with
aliquots of the cDNA bank wherein (i)
cotransfection with HLA-A*0201 DNA is carried
out or (ii) HLA-A*0201 positive recipient
cells are used,
 - (e) testing the transfected recipient cells for
their ability to stimulate T cells,
 - (f) identifying a cDNA sequence which codes for
the antigen which contains the peptide ligand
and

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(g) identifying the sequence of the peptide
ligand.

44. Method as claimed in claim 43,
w h e r e i n
step (e) comprises testing for the ability to lyse
TNF-sensitive cells.

add 4

add G1

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